54 GCC	108 TAC Y	162 TCC S	216 ACC T	270 CCC P	324 CTC L
GCA	ATC I	AAC N	ACG	ATG M	CAA Q
AGT	GCC	TGC	AAC N		AAG K
45 CCG	99 AAG K	153 TAC Y	207 CGG R	261 CCC P	315 ATC I
AGT	CCC P	AAG K	CCT	GAC D	GCC
AAA	CGG R	AGC S	CTG L	ATC I	
36 GTA	90 TAC Y	144 TTC F	198 GGC G	252 TCC S	306 CCT (P
AGT ACA	AGC	ATC I	ACC	GTC V	AGA R
AGT	TCC	GTA V	GCC	ATG M	GTG V
27 GAA	81 TCC S	135 AAG K		243 GCC A	297 AAA K
990	၁၅၅	CAG Q		GAC D	TAC
CGT	TCC	ATT I	TTC	GAC D	CCG
18 TGG	72 GGA G	126 CGC R	180 CTG L	234 ACC T	288 ACT T
299	ATG M	GGA G	GAC	ACC	
299	AGG AT	GAT D	ATG M	CTG L	GAA CGC E R
ე <u>ე</u> ე	252 CGC	117 ATC I	171 ATC I	225 CTG L	279 AAT TCA (N S)
သသ	999	GAC D	GAC D	TCC	AAT N
9 5' GCT CCC CGC GGC GGC	999 ၁၁9	CTG L	AGC GAC ATC S D I	ATC	GCG A
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FIGURE14

333 342 351 360 369 378
TCC GCT GGT GTC GAG GAC AGA ACC ACA AGC CGT GGC CAG TCT GCT GAG AGA
S A G V E D K R T T S R G Q S A E R

FIGUREIB

432 GCA A	486 CAG Q	540 GCG A	594 GAA E	648 AAA K	702 CTG	756 4AC 1
GGA		CTG	GCT	CTA	GAG E	GAT D
GAA E	TGC	GTG V	AAA K	GGA G	GAG E	TTG
423 AGG R		531 AGC S	585 CTG L	639 GAA E	693 AGG R	747 TTT F
CGG R		CAG Q	GAA E	TTG L	ATG M	AGT S
CCC	CCC	ATC I	AAT N	GAA E	AAG K	TAC
414 CAG Q	468 GAG E	522 TTA L	576 ATC I	630 GTG V	684 AAG K	738 AAG K
GAG	AGA R	GAA E	AAA K	CGC	ATT I	TGT
CTG	CCC	GAA E	TTC	AAA K	GAC D	CCC
405 GGC G	459 AGG R	513 AGA R	567 GCA A	621 GAG E	675 AGT S	729 TGC C
GTG V	CCC	GAG	AGA R	CTA L	AAG K	AAC N
GTT V	GAG E	CCA	TCA	GTC V	TGC	ACC
396 CGG R	450 GTA V	504 CCT P	558 TTC F	612 GCT A	666 AAA K	720 AGG R
AGA R	CAG Q			TTG L	GAG	AGC S
, GAC D	GGA G	CGC R	GAG E	CAC H	ATT I	AGC S
387 AGG R	441 AGT S		549 GCA A	603 AAT N	657 GAG E	711 AGA R
CCA CTG P L	GAA E	0 0 0	GTT V	GCA A	GTG V	GCC
CCA P	TTT	GAA	CAG Q	GTT V	GTG V	GCG

21. 1.

810 CTC	864 TGG W	918 CTG L	972 GTC V	1026 GTG V	1080 TCA S	.134 CAT H
CTG	CTT	999	${ m TGT}$	TGC C	1 TTC F	1 GAC D
TAC Y	TGG W	82 891 900 909 918 TG AGC TGC CTG GAG CAC ATG TAC CAC GAC CTC GGG CTG S C L E H M Y H D L G L	TTC	TTC	AAG K	CTG
801 AAG K	855 GTC V	909 GAC D	963 CTG L	.017 TGC C	071 GAG E	125 GAT D
CCC	GAC D	CAC H	TGG W	1 CAC H	1 CAG Q	CAC H
TAC Y	TTT F	TAC Y	AGG R	CGG R	CTC	TGC
792 ACT T	846 ACC T	900 ATG M	954 AGG R	008 TTC F	062 AGT S	116 ATC I
CCC	CCG P	CAC H	CTC	1 AAC N	1 TGC C	GCC A
GTT V	AAG K	GAG E	ACC	CAC H	CTC	GCG
783 GAT D	837 CGG R	891 CTG L	945 GTC V	999 TTC F	.053 TGG W	107 ACA T
CGC R	CTG	TGC	CCT P	CCC	1 GTC V	1 ATG M
CGA R	GCC	AGC	AAC N	AAC N	ATG M	CTA L
~ r \ ^.	828 GAG E	882 G CTG AGC ' L S (936 ATC I	990 AAC N	044 AGC S	098 ATC I
G ACT C	ATC I	ATG M	AGC S	AGA R	1 TAC Y	1 CTG L
E L	ACC	882 GAG ATG CTG A E M L S	TTC	TAC Y	ATG M	ATC I
765 AAG K	819 GAC E	873 AAT N	927 GAC D	981 AAC N	035 ATG M	1089 XG GAT A D
AAG K	CCA P	CCC	AGG R	GAC D	1 CAG Q	1 ACG T
765 CAC AAG AAG H K K	TCT CCA S P	GAG CCC E P	GTC V	CAC H	GCC	caa acg Q T

FIGUREID

1188 CGC	1242 ; ATC I	1296 AAG K	1350 CAT H	1404 AAC N	1458 TCT S	1512 GAG E
GTC V	CAG	TTC	AGA R	AGC	ATC	1 TTA L
GCG A	TTC	999 9	GCA A	TAC Y	GAT D	rta L
1179 CTG L	1233 GCC A	.287 GAT D	.341 ATG M	395 GAC D	449 TGT C	503 IGT (
GAG	GTG V	1 CCT P	1 GAC D	1 TTT F	TGC C	3AC '
ACA T	GCC	CCA P	ACT T	AAT N	AAA K	GTG (
1170 CGC R	1224 TGC C	1278 ATC I	332 GCC A	386 GAG E	440 ATA I	494 TGG (
GCC A	CAC	AAC N	1 TTG L	1 ATG M	1 TTG L	CCT P
AAT N	CAC H	TCC	ATC I	AAA K	ATT	GAG
1161 ATC I	1215 AAC N	1269 TTC F	.323 TTA L	.377 GAG E	431 ATG M	485 GCA A
CAG	GAG E	ATC I	1 ACA T	1 AAA K	1 AAG K	1 GTC
TAC Y	CTG L	AAC N	ATC I	TTC F	CTG	GAA
1152 ACG T	1206 CCG P	1260 TGC C	.314 ATG M	368 TCT S	422 CTG L	476 ATG M
AAC	TCA S	GAG E	GGA G	1 GAT D	ACC T	CCA P
AAC N	ATC I	CCT P	CAG Q	ATG M	ATG M	CGT R
1143 TAC Y	1197 GAC D	1251 GAG E	.305 CGA R	359 ATT I	413 CAC H	467 GTC V
	AAT N	GCC A	ATC I	1 GAA E	1 GAG E	1 GAG E
1143 1152 1161 1170 1179 1188 CCC GGC TAC AAC ACG TAC CAG ATC AAT GCC CGC ACA GAG CTG GCG GTC CGC P G Y N N T Y Q I N A R T E L A V R	TAC Y	CTC	CAG Q	GCA A	GAG E	AAC N

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1566 CCG P	1620 AAG K	1674 GAG E	1728 AAG K	.782 TCT S	836 GGA G	890 CCG
GCA	ATC	GTT V	CTG L	1 ACG T	1 GAA E	1 IGG
GTG V	TTC	ATG M	GAG E	TTG	AGT	, 29E
1557 CCT P	1611 GGG G	.665 CCC P	.719 GAG E	773 AGC S	827 AAC N	881 ACG (
CTT) ATT I	1 TTC F	1 TAC Y	1 GAC D	1 AAA K	18 rgg 7
299	CAG Q	CTC	CGC R	ACT T	GTG V	TTC '
1548 GAA E	1602 GCC A	1656 AAG K	.710 GAT D	.764 AAG K	818 GAT D	872 CAG '
TCA	ACA	ACC	CGA R	1 AAG K	1 AGA R	1 CTG
AAG K	GCC	GTG V	TCC	CAG Q	AGC S	TGG
1539 GAG E	1593 AAG K	1647 ACA T	701 GAA E	.755 TTA L	809 AGA R	863 GCG
CGT	ACC	GAA	TGG W	1 GAG E	1 GAG E	1 GGG
GAC D	GTG V	TTT F	CTT	AAA K	AGA R	990
1530 AGC S	1584 AAA K	1638 ATG M	1692 CCA P	.746 ATG M	.800 TCC S	854 AAG
CAG Q	GAC	CCA	CAG Q	GCC A	1 AAG K	1 GGA
ATG M	CGA R	ATC	CTG L	GAC D	GAG E	TGA
1521 TTT F	1575 GAC D	1629 CTG L	1683 ATG M	1737 GAT D	.791 ACC T	.845 GCC A
TAT Y	ATG M	GTC	ATC) ATA I	GCC A	1 TGT C
1521 GAA TAT TTT A E Y F M	TTC	1629 1638 1647 1656 1665 1674 TTT GTC CTG ATC CCA ATG TTT GAA ACA GTG ACC AAG CTC TTC CCC ATG GTT GAG F V L I P M F E T V T K L F P M V E	GAG E	CGG R	999 9	GAC D

FIGURE 1E

FIGURE 1F

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FIGURE 2A

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FIGURE 2B

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g829179 PDE9A PDE8A TA IA FA A L Σ SKERIKETLDPIDEVA NTPALEGVETPLEVGG G H 니 \mathcal{O} ΕC TDI TPL 0 F S 伍 K LEGV LOE S C 니 3 MV LL YF V L S S A D V L H A T A A A D V T O S T N CVAOMMY O F NPFHNFRH NPYHNSTH LH S Z Ή Ŀ Д Z 309 437 244

g829179 PDE9A PDE8A ICHDLDHPGYNNTYQINARTELAVRYNDISPLENHHCAV TIHDVDHPGRTNSFLCNAGSELAILYNDTAVLESHHAAL CIHDVDHPGLTNQFLVNSSSELALMYNDESVLENHHLAV R K 477

g829179 PDE9A PDE8A AEPE-CNIFSNIPPDGFKQIRQGMITLILATDMAR TGDDKCNIFKNMERNDYRTLRQGIIDMVLATEMTK QNQG-CDIFCNMQKKQRQTLRKMVIDIVLSTDMSK ILA AFOLT FKL O, ď Ø 389 517 324

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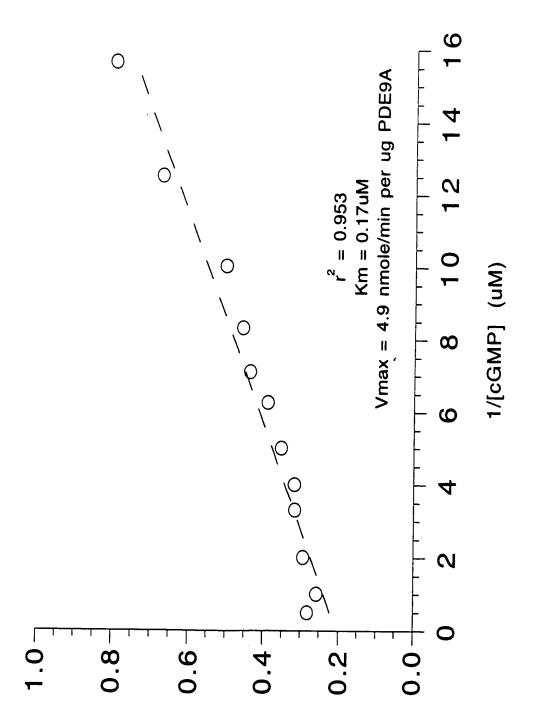
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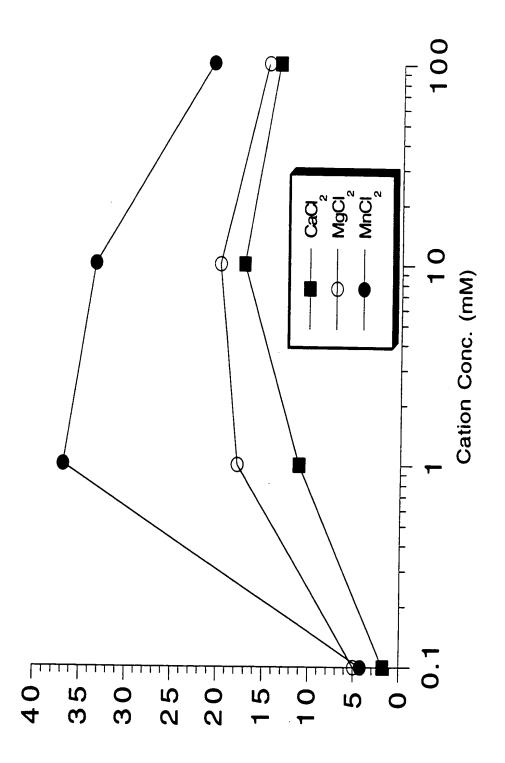
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FIGURE 2D



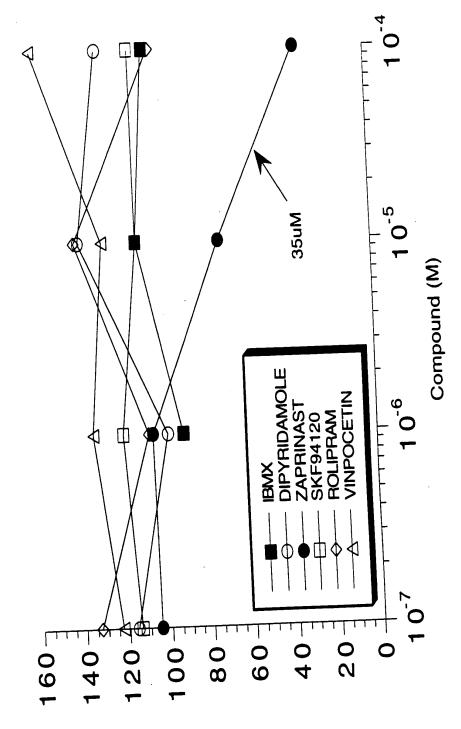
1/V (nmoles/min per ug PDE9A)



cGMP Hydrolysis (% conversion)

FIGURE 4

FIGURE 5



cGMP Hydrolysis (% control)